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PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2

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FULL ESTIMATED COST

CN

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STRUCTURE FILE UPDATES: 14 FEB 2002 HIGHEST RN 392654-43-2 DICTIONARY FILE UPDATES: 14 FEB 2002 HIGHEST RN 392654-43-2

TSCA INFORMATION NOW CURRENT THROUGH July 7, 2001

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

The P indicator for Preparations was not generated for all of the CAS Registry Numbers that were added to the H/Z/CA/CAplus files between 12/27/01 and 1/23/02. Use of the P indicator in online and SDI searches during this period, either directly appended to a CAS Registry Number or by qualifying an L-number with /P, may have yielded incomplete results. As of 1/23/02, the situation has been resolved. Also, note that searches conducted using the PREP role indicator were not affected.

Customers running searches and/or SDIs in the H/Z/CA/CAplus files incorporating CAS Registry Numbers with the P indicator between 12/27/01 and 1/23/02, are encouraged to re-run these strategies. Contact the CAS Help Desk at 1-800-848-6533 in North America or 1-614-447-3698, worldwide, or send an e-mail to help@cas.org for further assistance or to receive a credit for any duplicate searches.

```
=> s lidocaine/cn
            1 LIDOCAINE/CN
L1
=> d
L1
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS
     137-58-6 REGISTRY
RN
CN
     Acetamide, 2-(diethylamino)-N-(2,6-dimethylphenyl)- (9CI) (CA INDEX
NAME.)
OTHER CA INDEX NAMES:
     2',6'-Acetoxylidide, 2-(diethylamino)- (8CI)
OTHER NAMES:
     .alpha.-Diethylamino-2,6-acetoxylidide
CN
     2-(Diethylamino)-2',6'-acetoxylidide
CN
     2-(Diethylamino)-N-(2,6-dimethylphenyl)acetamide
CN
     Anbesol
CN
     Anestacon
CN
     Duncaine
CN
     Isicaina
CN
     Isicaine
CN
     Jetocaine
```

```
CN
     Lidocaine
CN
     Lignocaine
CN
     Maricaine
     Medicaine
CN
CN
     Remicaine
CN
     Rucaina
     Solcain
CN
CN
     Xilina
CN
     Xycaine
CN
    Xylestesin
CN
    Xyline
     Xylocain
CN
CN
    Xylocaine
CN
     Xylocitin
FS
     3D CONCORD
DR
     8059-42-5, 8059-66-3, 91484-71-8
MF
     C14 H22 N2 O
CI
     COM
LC
     STN Files:
                 ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS,
       CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DIOGENES,
       DRUGU, EMBASE, HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE,
       MRCK*, MSDS-OHS, NIOSHTIC, PHAR, PHARMASEARCH, PIRA, PROMT, RTECS*,
       SPECINFO, TOXCENTER, TOXLIT, USAN, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources: DSL**, EINECS**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
      NH-C-CH2-NEt2
Me
           Me
```

```
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
```

```
=> s morphine/cn
             1 MORPHINE/CN
L2
=> d
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2002 ACS
T.2
RN
     57-27-2 REGISTRY
     Morphinan-3, 6-diol, 7,8-didehydro-4,5-epoxy-17-methyl-
     (5.alpha., 6.alpha.) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Morphinan-3, 6.alpha.-diol, 7,8-didehydro-4,5.alpha.-epoxy-17-methyl-
CN
(8CI)
```

68 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

6147 REFERENCES IN FILE CA (1967 TO DATE)

6156 REFERENCES IN FILE CAPLUS (1967 TO DATE) 31 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
OTHER NAMES:
```

CN (-)-Morphine

CN Duromorph

CN 1-Morphine

CN Meconium

CN Morphia

CN Morphin

CN Morphina

CN Morphine

CN Morphinism

CN Morphinum

CN Morphium

CN Moscontin

CN Ospalivina

FS STEREOSEARCH

DR 8053-16-5, 85201-37-2, 47106-99-0

MF C17 H19 N O3

CI COM

LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,

BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DETHERM*, DIOGENES, DRUGU, EMBASE, GMELIN*, HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NAPRALERT, NIOSHTIC, PHAR, PHARMASEARCH,

PIRA,

PROMT, RTECS*, SPECINFO, TOXCENTER, TOXLIT, USAN, USPAT2, USPATFULL, VETU

(*File contains numerically searchable property data)
Other Sources: EINECS**

(**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry. Rotation (-).

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

17619 REFERENCES IN FILE CA (1967 TO DATE)

229 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

17630 REFERENCES IN FILE CAPLUS (1967 TO DATE)

=> file embase biosis medline embase uspatfull

COSȚ IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 11.48 11.63

FILE 'EMBASE' ENTERED AT 16:58:46 ON 15 FEB 2002

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FILE 'BIOSIS' ENTERED AT 16:58:46 ON 15 FEB 2002 COPYRIGHT (C) 2002 BIOLOGICAL ABSTRACTS INC.(R)

FILE 'MEDLINE' ENTERED AT 16:58:46 ON 15 FEB 2002

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FILE 'USPATFULL' ENTERED AT 16:58:46 ON 15 FEB 2002 CA INDEXING COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

=> file caplus COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL
ENTRY SESSION
3.29 14.92

FILE 'CAPLUS' ENTERED AT 16:58:52 ON 15 FEB 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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FILE COVERS 1907 - 15 Feb 2002 VOL 136 ISS 8 FILE LAST UPDATED: 14 Feb 2002 (20020214/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

The P indicator for Preparations was not generated for all of the CAS Registry Numbers that were added to the CAS files between 12/27/01 and 1/23/02. As of 1/23/02, the situation has been resolved. Searches and/or SDIs in the H/Z/CA/CAplus files incorporating CAS Registry Numbers with the P indicator executed between 12/27/01 and 1/23/02 may be incomplete. See the NEWS message on this topic for more information.

```
=> s lidocaine or 137-58-6/rn
7758 LIDOCAINE
6 LIDOCAINES
7758 LIDOCAINE
(LIDOCAINE OR LIDOCAINES)
6181 137-58-6
68 137-58-6D
6143 137-58-6/RN
(137-58-6 (NOTL) 137-58-6D)
L3 8957 LIDOCAINE OR 137-58-6/RN
```

```
=> s morphine or 57-27-2/rn
         34678 MORPHINE
           166 MORPHINES
         34725 MORPHINE
                  (MORPHINE OR MORPHINES)
         17796 57-27-2
           230 57-27-2D
         17667 57-27-2/RN
                  (57-27-2 (NOTL) 57-27-2D )
         35445 MORPHINE OR 57-27-2/RN
L4
\Rightarrow s 13 and 14
           416 L3 AND L4
=> s topical analgesic
         29361 TOPICAL
            30 TOPICALS
         29376 TOPICAL
                  (TOPICAL OR TOPICALS)
         32660 ANALGESIC
         27203 ANALGESICS
         40072 ANALGESIC
                  (ANALGESIC OR ANALGESICS)
L6
           110 TOPICAL ANALGESIC
                  (TOPICAL (W) ANALGESIC)
=> s 15 and 16
L7
            1 L5 AND L6
=> d 17
L7
     ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS
AN
     2000:779733 CAPLUS
DN
     134:66082
TΙ
     Analgesic synergy between topical lidocaine and topical opioids
ΑU
     Kolesnikov, Yuri A.; Chereshnev, Igor; Pasternak, Gavril W.
CS
     The Department of Anesthesiology, Memorial Sloan-Kettering Cancer Center,
     New York, NY, USA
SO
     J. Pharmacol. Exp. Ther. (2000), 295(2), 546-551
     CODEN: JPETAB; ISSN: 0022-3565
PB
     American Society for Pharmacology and Experimental Therapeutics
DT
     Journal
LA
     English
              THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 32
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
=> s 15 and topical
         29361 TOPICAL
            30 TOPICALS
         29376 TOPICAL
                  (TOPICAL OR TOPICALS)
L8
            19 L5 AND TOPICAL
=> s 18 and py<2000
      19714796 PY<2000
L9
            12 L8 AND PY<2000
=> dup rem 19
PROCESSING COMPLETED FOR L9
```

=> d l10 ab bib kwic

L10 ANSWER 1 OF 12 CAPLUS COPYRIGHT 2002 ACS

AB Methods and app. for improving dermal and mucosal administration of drugs through the use of controlled heat and other phys. means, i.e., ultrasound, microwave, elec. current, and vibrations, are described. The controlled heat and other phys. means are used to alter, mainly increase, the drug release rate from dermal drug delivery systems (DDDSs), conventional com. DDDSs, or drugs delivered into a sub-skin depot site

via

injection and other methods. For example, with heating by the temp. control app., it was found that fentanyl entered the systemic circulation of human volunteers earlier and at faster rate from a com. available dermal patch, Duragesic 50 (designed to deliver an av. of 50 g fentanyl/h), compared to the unheated patch. At 240 min, the end of the heating and fentanyl patch application, the av. serum concns. of fentanyl was about 5 times that of the unheated patch. These results demonstrates that controlled heat can significantly increase the speed of dermal fentanyl absorption and shorten the onset time. It is believed that the increased temp. increases the skin permeability resulting in the drug entering the patient's systemic circulation faster.

AN 2001:423696 CAPLUS

DN 135:37181

TI Methods and temperature control apparatus for improved administration of pharmaceutically active compounds including hormones

IN Zhang, Jie; Zhang, Hao

PA Zars, Inc., USA

SO U.S., 37 pp., Cont.-in-part of U.S. 5,919,479. CODEN: USXXAM

DT Patent

LA English

FAN.CNT 5

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PATENT NO.
                    KIND DATE
                                          APPLICATION NO. DATE
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                           _____
ΡI
                     В1
                            20010612
    US 6245347
                                          US 1998-162890
                                                          19980929
    US 5658583
                      Α
                            19970819
                                          US 1995-508463
                                                           19950728 <--
    WO 2000018339
                    A1
                            20000406
                                          WO 1999-US22698 19990929
            AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
            CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
             IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD,
            MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK,
            SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
            DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
            CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    AU 9964062
                      A1
                            20000417
                                          AU 1999-64062
                                                            19990929
    EP 1117357
                            20010725
                                          EP 1999-951669
                      Α1
                                                            19990929
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO
    US 6303142
                      В1
                            20011016
                                          US 2000-545591
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    US 6306431
                      В1
                            20011023
                                          US 2000-545497
                                                            20000407
    US 6340472
                      В1
                            20020122
                                          US 2000-545495
                                                            20000407
    US 2001037104
                      Α1
                                          US 2001-796250
                                                            20010228
                            20011101
PRAI US 1995-508463
                      A3
                           19950728
    US 1997-819880
                      A2
                           19970318
    US 1998-162890
                      Α
                           19980929
    US 1999-317313
                      Α
                           19990524
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Α
     US 1999-317372
                            19990524
     WO 1999-US22698 W
                            19990929
     US 2000-185913 P
                            20000229
     US 2000-545591
                       Α2
                            20000407
RE.CNT 38
              THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     PATENT NO.
                    KIND DATE
                                          APPLICATION NO. DATE
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     US 6245347 B1
                            20010612
PΙ
                                          US 1998-162890 19980929
     US 5658583
                     Α
                            19970819
                                          US 1995-508463
                                                            19950728 <--
                     A1 20000406
     WO 2000018339
                                          WO 1999-US22698 19990929
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             CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
             IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD,
             MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK,
             SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                      AU 1999-64062
     AU 9964062
                      A1
                            20000417
                                                          19990929
     EP 1117357
                                           EP 1999-951669 19990929
                      Α1
                            20010725
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                           US 2000-545591
     US 6303142
                     B1
                            20011016
                                                            20000407
     US 6306431
                      В1
                            20011023
                                           US 2000-545497
                                                            20000407
     US 6340472
                      В1
                            20020122
                                           US 2000-545495
                                                            20000407
     US 2001037104
                     A1
                            20011101
                                           US 2001-796250
                                                            20010228
ΙT
     Drug delivery systems
        (topical, mucosal; controlled heat and other phys. means for
        improved dermal and mucosal drug delivery)
IT
     50-02-2, Dexamethasone 50-28-2, Estradiol, biological studies
50-81-7,
     Vitamin C, biological studies
                                   51-55-8, Atropine, biological studies
     54-11-5, Nicotine 55-63-0, Nitroglycerin 57-27-2,
    Morphine, biological studies 57-42-1, Meperidine
                                                          57-83-0.
     Progesterone, biological studies 58-20-8, Testosterone cypionate
     58-22-0, Androderm
                        94-24-6, Tetracaine 113-15-5, Ergotamine
     137-58-6, Lidocaine
                         315-37-7, Testosterone enanthate
     511-12-6, Dihydroergotamine 721-50-6, Prilocaine
                                                         1406-18-4, Vitamin E
     4205-90-7, Clonidine 9002-89-5, Polyvinyl alcohol 9004-10-8, Insulin,
     biological studies 11103-57-4, Vitamin A 26780-50-7, Medisorb 8515DL
     34346-01-5, Atrigel 38396-39-3, Bupivacaine 56030-54-7, Sufentanil
     71195-58-9, Alfentanil 103628-46-2, Sumatriptan 131723-69-8, Smart
     Hydrogel 132875-61-7, Remifentanil 139264-17-8, Zolmitriptan
     144034-80-0, Rizatriptan
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (controlled heat and other phys. means for improved dermal and mucosal
        drug delivery)
```

=> d 110 2-12 ab bib kwic

L10 ANSWER 2 OF 12 CAPLUS COPYRIGHT 2002 ACS

AB A method for enhancing the analgesic efficacy of opioids and local anesthetics that act on peripheral sensory nerves in tissues, and preferably in non-inflamed tissue, in a subject in need of such treatment,

is based on administration of a therapeutically effective amt. of an opioid, opioid peptide (derivs. of naturally occurring endorphins), a

local anesthetic or mixts. thereof in combination with a hyperosmolar soln. having an osmolality of > 300 mOsm/L and optionally other pharmaceutically acceptable carriers and diluents. The method is applicable in a large variety of painful conditions, such as injury of skin (burns, radiation, cuts, psoriasis, surgery, infections, etc.), cancer, musculocutaneous and myofascial pain syndromes, causalgia, shingles, postherpetic neuralgia, headache, and gastrointestinal, facial, urol., abdominal gynecol. or postoperative pain. The advantage of the method is pain relief by using extremely small doses of the active agent in combination with hyperosmolar solns. and thereby foregoing all the untoward systemic side effects of opiates or local anesthetics. The antinociceptive effects in noninflamed and inflamed rat paws after concomitant injection of hyperosmolar soln. of mannitol (1M) with three opioid agonists, DAGO and DPDPE (0.004 mg) or U-50488H (0.04 mg) were examd. All three opioid agonists, in combination with mannitol, produced elevations in paw pressure threshold in noninflamed paws that were comparable to those of inflamed paws at 12 h and 4 days after inoculation. The addn. of mannitol to opioids did not alter their antinociceptive effects in inflamed paws. 1999:571725 CAPLUS 131:194297 Method of enhancing the analgesic efficacy of locally and topically administered opioids and other local anesthetics Stein, Christoph El Khoury & Stein, Ltd., USA U.S., 11 pp., Cont. of U.S. Ser. No. 488,021, abandoned. CODEN: USXXAM Patent English FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE US 5948389 A 19990907 US 1997-922573 19970903 <--PRAI US 1995-488021 19950607 THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT US 5948389 A **19990907** US 5948389 - AIND DATE APPLICATION NO. DATE US 5948389 A 19990907 US 1997-922573 19970903 <--Drug delivery systems (topical; hyperosmolar solns. for enhancing analgesic efficacy of locally and topically administered opioids and local anesthetics) **57-27-2**, biological studies 57-42-1, Meperidine 69-65-8, D-Mannitol 94-24-6, Pontocaine 103-81-1, Benzeneacetamide **137-58-6, Lidocaine** 721-50-6, Prilocaine 990-73-8, Fentanyl citrate 3572-80-3, Cyclazocine 36637-18-0, Etidocaine 38396-39-3, Bupivacaine 74135-04-9, Morphiceptin 78123-71-4, DAGO 83386-35-0, Tifluadom 83913-06-8, U 50488H 88373-73-3 RL: BAC (Biological activity or effector, except adverse); THU

T.10 ANSWER 3 OF 12 CAPLUS COPYRIGHT 2002 ACS

the

ΑN

DN

IN

PA

SO

DT

LA

PΙ

PΙ

ΙT

AB We prospectively studied topical lidocaine-prilocaine cream (EMLA) vs. IV morphine in a double-blinded, randomized

(Therapeutic use); BIOL (Biological study); USES (Uses)

topically administered opioids and local anesthetics)

(hyperosmolar solns. for enhancing analgesic efficacy of locally and

```
fashion for pain relief during thoracostomy tube (chest tube; CT)
removal.
     Adult patients who had undergone thoracotomy or median sternotomy were
     randomized to receive either EMLA cream over CT sites transdermally for 3
     h or IV morphine 0.5 h before CT removal. Pain behavior was
     obsd. and rated before, during, and after CT removal. Pain behavior
     increased less in the topical EMLA group (mean .+-. SE,
     4.4.+-.0.39) compared with the IV morphine group (6.0.+-.0.38; P
     < 0.01). No signs of infection were noted at the CT sites 24 or 48 h
     after CT removal. We conclude that EMLA cream is more effective than IV
     morphine in preventing the pain assocd. with CT removal.
     Implications: Postoperatively applying a topical anesthetic
     cream onto chest tube sites of chest surgery patients 3 h before chest
     tube removal is more effective than IV morphine in blunting pain
     response.
     1999:319708 CAPLUS
ΑN
DN
     130:347345
ΤI
     Topical lidocaine-prilocaine cream (EMLA) for
     thoracostomy tube removal
ΑU
     Valenzuela, Roberto C.; Rosen, David A.
CS
     Department of Anesthesiology, West Virginia University, Morgantown, WV,
     26506-9134, USA
     Anesth. Analg. (Baltimore) (1999), 88(5), 1107-1108
SO
     CODEN: AACRAT; ISSN: 0003-2999
PB
     Lippincott Williams & Wilkins
DT
     Journal
LA
     English
RE.CNT 6
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
ΤI
     Topical lidocaine-prilocaine cream (EMLA) for
     thoracostomy tube removal
SO
     Anesth. Analg. (Baltimore) (1999), 88(5), 1107-1108
     CODEN: AACRAT; ISSN: 0003-2999
AΒ
     We prospectively studied topical lidocaine-prilocaine
     cream (EMLA) vs. IV morphine in a double-blinded, randomized
     fashion for pain relief during thoracostomy tube (chest tube; CT)
removal.
     Adult patients who had undergone thoracotomy or median sternotomy were
     randomized to receive either EMLA cream over CT sites transdermally for 3
     h or IV morphine 0.5 h before CT removal. Pain behavior was
     obsd. and rated before, during, and after CT removal. Pain behavior
     increased less in the topical EMLA group (mean .+-. SE,
     4.4.+-.0.39) compared with the IV morphine group (6.0.+-.0.38; P
     < 0.01). No signs of infection were noted at the CT sites 24 or 48 h
     after CT removal. We conclude that EMLA cream is more effective than IV
     morphine in preventing the pain assocd. with CT removal.
     Implications: Postoperatively applying a topical anesthetic
     cream onto chest tube sites of chest surgery patients 3 h before chest
     tube removal is more effective than IV morphine in blunting pain
     response.
ST
     lidocaine prilocaine cream anesthetic analgesic thoracostomy
TΤ
     Topical drug delivery systems
        (anesthetics; topical lidocaine-prilocaine cream
        (EMLA) vs. i.m. morphine effect on thoracostomy tube removal
        in humans)
ŢΤ
     Surgery
        (thoracostomy tube; topical lidocaine-prilocaine
        cream (EMLA) vs. i.m. morphine effect on thoracostomy tube
        removal in humans)
ΙT
    Analgesics
```

```
(topical lidocaine-prilocaine cream (EMLA) vs. i.m.
        morphine effect on thoracostomy tube removal in humans)
IT
     Anesthetics
        (topical; topical lidocaine-prilocaine
        cream (EMLA) vs. i.m. morphine effect on thoracostomy tube
        removal in humans)
                            721-50-6, Prilocaine
IT
     137-58-6, Lidocaine
     101362-25-8, EMLA
     RL: BAC (Biological activity or effector, except adverse); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (topical lidocaine-prilocaine cream (EMLA) vs. i.m.
        morphine effect on thoracostomy tube removal in humans)
    ANSWER 4 OF 12 CAPLUS COPYRIGHT 2002 ACS A pharmaceutical compn. contains an acid addn. salt of a basic drug and a
L10
     fatty acid or bile acid. The acid addn. salts thus formed exhibit
     enhanced transmucosal and transdermal penetration of the basic drug.
     acid addn. salts, an inclusion complex contg. said salts and a use of
said
     salts are also disclosed. Thus, 10.01 g capric acid was added to a soln.
     of 13.91 g salbutamol base in 600 mL ethanol and stirred until all solid
     material was dissolved. Evapn. of the solvent gave a pale yellow oil to
     which 300 mL warm Et acetate was added and stored at 5.degree. for 36 h
     resulting in the pptn. of a fine white solid which was sepd. and purified
     to obtain salbutamol caprate. A sublingual table contained salbutamol
     caprate .gamma.-cyclodextrin complex (prepn. given) 32, lactose 20, and
     magnesium stearate 1 mg.
ΑN
     1998:66112 CAPLUS
DN
     128:145353
ΤI
     Pharmaceutical composition containing acid addition salt of basic drug
     Penkler, Lawrence John; De Kock, Lueta-Ann; Whittaker, Darryl Vanstone
ΙN
PA
     Farmarc Nederland B.V., Neth.; Dyer, Alison Margaret; Penkler, Lawrence
     John; De Kock, Lueta-Ann; Whittaker, Darryl Vanstone
SO
     PCT Int. Appl., 51 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 2
     PATENT NO.
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                             DATE
                                            APPLICATION NO. DATE
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JP 2001508027

T2

20010619

JP 1998-505726

19970711

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                                           US 1999-225470
PRAI ZA 1996-5889
                      A
                            19960711
     WO 1997-GB1873
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                           19970711
PΙ
     WO 9802187 A1 19980122
     PATENT NO.
                    KIND DATE
                                         APPLICATION NO. DATE
                     A1 19980122 WO 1997-GB1873 19970711 <--
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             DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ,
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             PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US,
             UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG
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                                          EP 1997-930681 19970711
        R: AT, BE, DE, ES, FR, GB, IT
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                            20000425
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                            20000425
                                           KR 1999-700167
                                                           19990111
     US 6255502
                      В1
                            20010703
                                           US 1999-225470
                                                          19990419
ΙT
     Suppositories (drug delivery systems)
     Tablets (drug delivery systems)
       Topical gels (drug delivery systems)
     Transdermal drug delivery systems
        (pharmaceutical compn. contq. acid addn. salt of basic drug)
ΙT
     52-53-9, Verapamil 57-27-2, Morphine, biological
             59-46-1, Procaine 82-92-8, Cyclizine 137-58-6,
     Lidocaine 364-62-5, Metoclopramide 437-38-7, Fentanyl
     915-30-0, Diphenoxylate 7585-39-9D, .beta.-Cyclodextrin, hydroxypropyl
     derivs. 12619-70-4, Cyclodextrin 12619-70-4D, Cyclodextrin, complexes
     13392-18-2, Fenoterol 17465-86-0, .gamma.-Cyclodextrin
                                                                23031-25-6,
     Terbutaline 38396-39-3, Bupivacaine 53179-11-6, Loperamide
     55985-32-5, Nicardipine 62571-86-2, Captopril 74913-18-1, Dynorphin
     75847-73-3, Enalapril 87333-19-5, Ramipril 89365-50-4, Salmeterol
     103628-46-2, Sumatriptan 121679-13-8, Naratriptan 139264-17-8,
     Zolmitriptan 144034-80-0, Rizatriptan 202282-68-6 202282-69-7
     202282-70-0
                  202282-71-1 202282-74-4
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmaceutical compn. contg. acid addn. salt of basic drug)
L10
    ANSWER 5 OF 12 CAPLUS COPYRIGHT 2002 ACS
AB
     4-Decyl-2-oxazolidinone (SR 38) formulated in transdermal delivery system
     contg. lipophilic and hydrophilic drugs enhanced the drug permeation
     through the human stratum corneum. In addn., the enhanced the drug
     permeation from topical formulations. There was no skin
     sensitization or irritation.
AN
     1997:463797 CAPLUS
DN
     127:113267
TI
     Oxazolidinones: a new class of cyclic urethane transdermal enhancer
(CUTE)
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ΑU
     Pfister, William R.; Rajadhyaksha, Vithal J.
CS
     Pharmaceutical Research and Development, Pharmetrix Div. of TCPI, Menlo
     Park, CA, 94025, USA
SO
     Proc. Int. Symp. Controlled Release Bioact. Mater. (1997), 24th,
     709-710
     CODEN: PCRMEY; ISSN: 1022-0178
PΒ
     Controlled Release Society, Inc.
     Journal
DT
     English
LA
     Proc. Int. Symp. Controlled Release Bioact. Mater. (1997), 24th,
SO
     709-710
     CODEN: PCRMEY; ISSN: 1022-0178
AB
     4-Decyl-2-oxazolidinone (SR 38) formulated in transdermal delivery system
     contg. lipophilic and hydrophilic drugs enhanced the drug permeation
     through the human stratum corneum. In addn., the enhanced the drug
     permeation from topical formulations. There was no skin
     sensitization or irritation.
ST
     oxazolidinone SR 38 transdermal drug enhancer; topical drug
     enhancer SR 38
     Permeation (biological)
TΤ
     Skin
       Topical drug delivery systems
     Transdermal drug delivery systems
        (oxazolidinone as cyclic urethane transdermal enhancer)
ΙT
     50-23-7, Hydrocortisone 53-86-1, Indomethacin
                                                       57-83-0, Progesterone,
     biological studies 64-31-3, Morphine sulfate
                                                      87 - 33 - 2,
     Isosorbide dinitrate 137-58-6, Lidocaine
                                                721-50-6,
                 15307-86-5, Diclofenac
     Prilocaine
                                          21829-25-4, Nifedipine
28981-97-7,
     Alprazolam
                  33286-22-5, Diltiazem hydrochloride
                                                      38304-91-5, Minoxidil
     RL: BPR (Biological process); THU (Therapeutic use); BIOL (Biological
     study); PROC (Process); USES (Uses)
        (oxazolidinone as cyclic urethane transdermal enhancer)
L10
    ANSWER 6 OF 12 CAPLUS COPYRIGHT 2002 ACS
     A review with many refs. identifying and analyzing the controlled clin.
     trial data for peripheral neuropathic pain (PNP) and complex regional
pain
     syndromes (CRPS). A total of 72 articles were found, which included 92
     controlled drug trials using 48 different treatments. The methods of
     these studies were critically reviewed and the results summarized and
     compared. The PNP trial literature gave consistent support (two or more
     trials) for the analgesic effectiveness of tricyclic antidepressants,
i.v.
     and topical lidocaine, i.v. ketamine, carbamazepine
     and topical aspirin. There was limited support (one trial) for
     the analgesic effectiveness of oral, topical and epidural
     clonidine and for s.c. ketamine. The trial data were contradictory for
    mexiletine, phenytoin, topical capsaicin, oral non-steroidal
     anti-inflammatory medication, and i.v. morphine. Anal. of the
     trial methods indicated that mexiletine and i.v. morphine were
     probably effective analgesics for PNP, while non-steroidal agents were
    probably ineffective. Codeine, magnesium chloride, propranolol,
    lorazepam, and i.v. phentolamine all failed to provide analgesia in
```

trials. There were no long-term data supporting the analgesic effectiveness of any drug and the etiol. of the neuropathy did not predict treatment outcome. Review of the controlled trial literature for CRPS

sinale

identified several potential problems with current clin. practices. The

trial data only gave consistent support for analgesia with corticosteroids, which had long-term effectiveness. There was limited support for the analgesic effectiveness of topical dimethylsulfoxide (DMSO), epidural clonidine and i.v. regional blocks (IVRBs) with bretylium and ketanserin. The trial data were contradictory for intranasal calcitonin and i.v. phentolamine and anal. of the trial methods indicated that both treatments were probably ineffective for most patients. There were consistent trial data indicating that quanethidine and reserpine IVRBs were ineffective, and limited trial data indicating that droperidol and atropine IVRBs were ineffective. No placebo controlled data were available to evaluated sympathetic ganglion blocks (SGBs) with local anesthetics, surgical sympathectomy, or phys. therapy. Only the capsaicin trials presented data which allowed for meta-anal. This meta-anal. demonstrated a significant capsaicin effect with a pooled odds ratio of 2.35 (95 confidence intervals 1.48, 3.22). The methods scores were higher for the PNP trials (66.2) than the CRPS trials (57.6). The CRPS trials tended to use less subjects and were less likely to use placebo controls, double-blinding, or perform statistical tests for differences in outcome measures between groups. There was almost no overlap in the controlled trial literature between treatments for PNP and CRPS, and treatments used in both conditions (i.v. phentolamine and epidural clonidine) had similar results.

AN 1997:723920 CAPLUS

DN 128:123374

TI A critical review of controlled clinical trials for peripheral neuropathic

pain and complex regional pain syndromes

AU Kingery, Wade S.

CS Miranda Ave., Physical Medicine and Rehabilitation Service (117), Veterans

Affairs Palo Alto Health Care System, Palo Alto, CA 94304, 3801, USA SO Pain (1997), 73(2), 123-139 CODEN: PAINDB; ISSN: 0304-3959

PB Elsevier

DT Journal; General Review

LA English

SO Pain (1997), 73(2), 123-139 CODEN: PAINDB; ISSN: 0304-3959

AB A review with many refs. identifying and analyzing the controlled clin. trial data for peripheral neuropathic pain (PNP) and complex regional pain

syndromes (CRPS). A total of 72 articles were found, which included 92 controlled drug trials using 48 different treatments. The methods of these studies were critically reviewed and the results summarized and compared. The PNP trial literature gave consistent support (two or more trials) for the analgesic effectiveness of tricyclic antidepressants,

i.v.

and topical lidocaine, i.v. ketamine, carbamazepine and topical aspirin. There was limited support (one trial) for the analgesic effectiveness of oral, topical and epidural clonidine and for s.c. ketamine. The trial data were contradictory for mexiletine, phenytoin, topical capsaicin, oral non-steroidal anti-inflammatory medication, and i.v. morphine. Anal. of the trial methods indicated that mexiletine and i.v. morphine were probably effective analgesics for PNP, while non-steroidal agents were probably ineffective. Codeine, magnesium chloride, propranolol, lorazepam, and i.v. phentolamine all failed to provide analgesia in single

trials. There were no long-term data supporting the analgesic effectiveness of any drug and the etiol. of the neuropathy did not predict

treatment outcome. Review of the controlled trial literature for CRPS identified several potential problems with current clin. practices. The trial data only gave consistent support for analgesia with corticosteroids, which had long-term effectiveness. There was limited support for the analgesic effectiveness of topical dimethylsulfoxide (DMSO), epidural clonidine and i.v. regional blocks (IVRBs) with bretylium and ketanserin. The trial data were contradictory for intranasal calcitonin and i.v. phentolamine and anal. of the trial methods indicated that both treatments were probably ineffective for most patients. There were consistent trial data indicating that quanethidine and reserpine IVRBs were ineffective, and limited trial data indicating that droperidol and atropine IVRBs were ineffective. No placebo controlled data were available to evaluated sympathetic ganglion blocks (SGBs) with local anesthetics, surgical sympathectomy, or phys. therapy. Only the capsaicin trials presented data which allowed for meta-anal. This meta-anal. demonstrated a significant capsaicin effect with a pooled odds ratio of 2.35 (95 confidence intervals 1.48, 3.22). The methods scores were higher for the PNP trials (66.2) than the CRPS trials (57.6). The CRPS trials tended to use less subjects and were less likely to use placebo controls, double-blinding, or perform statistical tests for differences in outcome measures between groups. There was almost no overlap in the controlled trial literature between treatments for PNP and CRPS, and treatments used in both conditions (i.v. phentolamine and epidural clonidine) had similar results.

L10 ANSWER 7 OF 12 CAPLUS COPYRIGHT 2002 ACS

AB Unit doses of a drug in cream, emulsion, gel, suspension, or ointment form

are placed in sep. compartments of a carrier for use in transdermal therapy. Thus, a blister pack contained 28 .apprx.1-mL numbered transparent compartments, of which compartments 1-14 each contained 3 mg 17.beta.-estradiol in 1 mL vehicle and compartments 15-28 each contained

mg 17.beta.-estradiol and 1 mg norethisterone acetate in 1 mL vehicle. The compartments had an air-tight seal of Al foil, which could be removed sep. for each compartment for daily application of a unit dose to the skin

for treatment of postmenopausal symptoms.

AN 1994:144205 CAPLUS

DN 120:144205

TI Unit doses of a semisolid topical drug for transdermal therapy

IN Liedtke, Rainer K.

PA Germany

3

SO Ger. Offen., 4 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 1

CNII				
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 4223004	A1	19940120	DE 1992-4223004	19920713 <
EP 581057	A1	19940202	EP 1993-110690	19930705 <
EP 581057	B1	19981007		
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AT 171868	E	19981015	AT 1993-110690	19930705 <
ES 2123595	Т3	19990116	ES 1993-110690	19930705 <
JP 07275321	A2	19951024	JP 1993-172931	19930713 <
US 5686112	A	19971111	US 1995-569958	19951220 <
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ΤI
     Unit doses of a semisolid topical drug for transdermal therapy
     DE 4223004 A1 19940120
PΙ
    DE 4223004 A1 19940120 DE 1992-4223004 19920713 <--
EP 581057 A1 19940202 EP 1993-110690 19930705 <--
EP 581057 B1 19981007
R: AT, BE, CH DE DE CO
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US 5686112 A 19971111 US 1995-569958 19951220 <--
ΙT
     50-56-6, Oxytocin, biological studies 54-11-5, Nicotine 57-27-2
     , Morphine, biological studies 57-27-2D, Morphine,
     derivs. 96-88-8, Mepivacaine 136-47-0, Pantocaine 137-58-6,
     Lidocaine 342-10-9, Kallidin 437-38-7, Fentanyl 721-50-6,
     Prilocaine 8011-61-8, Tyrocidine 9002-72-6, Growth hormone
     9004-10-8, Insulin, biological studies 9007-12-9, Calcitonin
     9011-97-6, Cholecystokinin 9034-40-6D, LHRH, analogs 9061-61-4, Nerve
     growth factor 11096-26-7, Erythropoietin 16679-58-6, Desmopressin
     52485-79-7, Buprenorphine 85637-73-6, Atrial natriuretic factor
     143011-72-7, G-CSF
     RL: BIOL (Biological study)
        (kit contg. semisolid transdermal unit doses of)
L10 ANSWER 8 OF 12 CAPLUS COPYRIGHT 2002 ACS
AΒ
     The title gels have improved thermorheolog, properties and a gelling
temp.
     interval of approx. 25-37.degree.; the gels comprise (1) 10-30 wt.% of
     block copolymers of
.alpha.-hydro-.omega.-hydroxypoly(oxyethylene)/poly(ox
     ypropylene)/poly(oxyethylene) (Poloxamer)
H(OCH2CH2) a [OCH(CH3) CH2]b(OCH2CH
     2) aOH (a .gtoreq.2; b .gtoreq.15; total proportion of hydrophilic
     polyethylene units is 20-90 wt.% of the copolymer having a mol. wt. of
     1000-16,000); (2) 0.01-5 wt.% carboxyvinyl polymer (Carbomer) of mol. wt.
     1 \times 106-4 \times 106; (3) sufficient pharmaceutically acceptable base to
adjust
     the pH to 4-8; (4) 20-85 wt.% water; and (5) optional usual auxiliary
     agents. The liq. formulations may be used for .beta.-lactam antibiotics,
     antibacterials, chemotherapeutics, antiinflammatories, cosmetics, etc. A
     liq. thermoreversible formulation of betamethasone-17,21-dipropionate (I)
     contained I 0.05, Pluronic F127 18.0, Carbopol 934P 0.3, 10%ag. NaOH 5,
     and demineralized water to 100 wt.%.
ΑN
     1993:567753 CAPLUS
DN
     119:167753
TI
     Thermoreversible gel as a liquid pharmaceutical carrier for a galenic
ΙN
     Kramaric, Anton; Resman, Aleksander; Kofler, Bojan; Zmitek, Janko
     LEK, Tovarna Farmacevtskih in Kemicnih Izdelkov, d.d., Slovenia
PΑ
SO
     Eur. Pat. Appl., 23 pp.
     CODEN: EPXXDW
DT
     Patent
T.A
     English
FAN.CNT 1
    PATENT NO. KIND DATE APPLICATION NO. DATE
     EP 551626 A1 19930721 EP 1992-121410 19921216 <--
PΙ
       R: AT, DE, FR, GB, IT, NL
     JP 05262670 A2 19931012 JP 1992-338663 19921218 <--
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PRAI YU 1991-17
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     EP 551626 A1 19930721
     PATENT NO. KIND DATE
                                         APPLICATION NO. DATE
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     EP 551626 A1 19930721
PΙ
                                          EP 1992-121410 19921216 <--
         R: AT, DE, FR, GB, IT, NL
     JP 05262670 A2 19931012
                                          JP 1992-338663 19921218 <--
IΤ
     Anesthetics
        (topical, thermoreversible gel carrier contg. Poloxamer and
        Carbomer for)
ΙT
     Pharmaceutical dosage forms
        (topical, thermoreversible gel carrier for, Poloxamer and
        carbomer in)
     50-02-2, Dexamethasone 50-36-2, Cocaine 50-78-2, Acetylsalicylic acid
IT
     51-34-3, Scopolamine 51-43-4, Epinephrine 51-55-8, Atropine,
     biological studies 53-86-1, Indomethacin 54-42-2, Idoxuridine
     57-27-2, uses 57-47-6, Physostigmine 58-71-9 59-46-1,
     Procaine 59-99-4, Neostigmine 69-53-4, Ampicillin 73-78-9, Lidocaine hydrochloride 79-57-2, Oxytetracycline 87-33-2,
     Isosorbide dinitrate 92-13-7, Pilocarpine 94-36-0, Benzoyl peroxide,
     biological studies 96-88-8, Mepivacaine 137-58-6,
     Lidocaine 154-21-2, Lincomycin 299-42-3, Ephedrine
     Betamethasone 530-43-8, Chloramphenicol palmitate 564-25-0,
     Doxycycline 1400-61-9, Nystatin 1404-90-6, Vancomycin 2180-92-9, Bupivacaine 2392-39-4, Dexamethasone-21-phosphate disodium salt
     5104-49-4, Flurbiprofen 5536-17-4 5593-20-4, Betamethasone-17,21-
     dipropionate 7553-56-2, Iodine, biological studies 9004-10-8,
Insulin,
                        9007-12-9, Calcitonin 9039-53-6, Urokinase
     biological studies
     11096-26-7, Erythropoietin 15307-79-6, Diclofenac sodium 15686-71-2,
    Cephalexin 15687-27-1, Ibuprofen 15826-37-6 16051-77-7, Isosorbide mononitrate 18323-44-9, Clindamycin 21829-25-4, Nifedipine
     22071-15-4, Ketoprofen 22199-08-2, Silver sulfadiazine 22494-42-4,
     Diflunisal 22832-87-7, Miconazole nitrate 23593-75-1, Clotrimazole
     24169-02-6, Econazole nitrate 24729-96-2, Clindamycin-2-phosphate
     26787-78-0, Amoxicillin 26839-76-9 26921-17-5, Timolol maleate
     34580-14-8, Ketotifen fumarate 35607-66-0, Cefoxitin 36322-90-4,
     Piroxicam 38194-50-2, Sulindac 38304-91-5, Minoxidil 39562-70-4,
     Nitrendipine 41340-25-4, Etodolac 42924-53-8, Nabumetone
51940-44-4,
     Pipemidic acid 52485-79-7, Buprenorphine 53648-05-8, Ibuproxam
     53994-73-3, Cefaclor 54527-84-3, Nicardipine hydrochloride
59277-89-3,
     Acyclovir 59995-64-1D, Thienamycin, derivs.
                                                   60607-34-3, Oxatomide
     60731-46-6, Elcatonin 64221-86-9 64485-93-4, Cefotaxime sodium
     64544-07-6, Cefuroxime axetil 65141-46-0, Nicorandil
                                                             65277-42-1,
     Ketoconazole
                  66085-59-4, Nimodipine 68401-82-1, Ceftizoxime sodium
     69049-73-6, Nedocromil 70458-96-7, Norfloxacin 72509-76-3, Felodipine
     74103-06-3, Ketorolac 74578-69-1, Ceftriaxone disodium 75695-93-1,
     Isradipine 79307-93-0, Azelastine hydrochloride 79660-72-3,
Fleroxacin
     82410-32-0, Gancyclovir 82419-36-1, Ofloxacin 85721-33-1,
     Ciprofloxacin 91524-15-1 93106-60-6, Enrofloxacin 111470-99-6
     114394-67-1 139639-23-9, Tissue plasminogen activator 150106-85-7
     150106-86-8
     RL: BIOL (Biological study)
        (dosage forms of, thermoreversible gel carrier contg. Poloxamer and
        Carbomer for)
```

AB Artificial composite membranes composed of silicone and pHEMA [poly(2-hydroxyethyl methacrylate] were developed as an alternative for skin membranes. The structure of the composite membranes was designed based on a model simulation of drug permeation properties. Composite membrane permeabilities for a wide range of drugs with diverse physicochem. properties were measured and compared with those of excised hairless rat skin. A reasonable correlation was found between the calcd. and obsd. permeability coeffs., and between the obsd. values for the composite and excised skin membranes. It is suggested that human skin permeability of drugs may be predicted by using slightly modified composite membranes.

AN 1992:67060 CAPLUS

DN 116:67060

TI Prediction of skin permeability of drugs. II. Development of composite membrane as a skin alternative

AU Hatanaka, Tomomi; Inuma, Masami; Sugibayashi, Kenji; Morimoto, Yasunori

CS Fac. Pharm. Sci., Josai Univ., Sakado, 350-02, Japan

SO Int. J. Pharm. (1992), 79(1), 21-8 CODEN: IJPHDE; ISSN: 0378-5173

DT Journal

LA English

SO Int. J. Pharm. (1992), 79(1), 21-8 CODEN: IJPHDE; ISSN: 0378-5173

IT Pharmaceutical dosage forms

(topical, drug permeation through composite membrane as alternative for human skin in relation to)

IT 50-28-2, .beta.-Estradiol, properties 51-21-8, 5-Fluorouracil 51-30-9,

Isoproterenol hydrochloride 52-26-6, **Morphine** hydrochloride 52-31-3, Cyclobarbital 53-86-1, Indomethacin 58-15-1, Aminopyrine 59-92-7, Levodopa, properties 60-80-0, Antipyrine 62-31-7, Dopamine hydrochloride 87-33-2, Isosorbide dinitrate **137-58-6**,

Lidocaine 5104-49-4, Flurbiprofen 15307-79-6, Diclofenac sodium 15687-27-1, Ibuprofen 22071-15-4, Ketoprofen 65141-46-0, Nicorandil

RL: PRP (Properties)

(permeation of, through composite membrane, as alternative for human skin)

L10 ANSWER 10 OF 12 CAPLUS COPYRIGHT 2002 ACS

AB To measure the contribution of lipid and pore (aq.) pathways to the total skin permeation of drugs, and to establish a predictive method for the steady state permeation rate of drugs, the relationship between permeability through excised hairless rat skin and some physicochem. properties of several drugs were compared with those through polydimethyl siloxane (silicone) and poly(2-hydroxyethyl methacrylate) (pHEMA) membranes, as typical soln.-diffusion and porous membranes, resp. A linear relation was found between the permeability coeffs. of drugs for the silicone membrane and their octanol/water partition coeffs. For the pHEMA membrane, the permeability coeffs. were almost const. independent of

the partition coeff. On the other hand, the skin permeation properties were classified into 2 types: one involves the case of lipophilic drugs, where the permeability coeff. is correlated to the partition coeff., similar to the silicone membrane; and the other involves hydrophilic drugs, where the permeability coeffs. were almost const., similar to pHEMA

membrane. The stratum corneum, the main barrier in skin, could be described as a membrane having 2 parallel permeation pathways: lipid and pore pathways. An equation for predicting the steady state permeation

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ΑN
     1991:128931 CAPLUS
DN
     114:128931
ΤI
     Prediction of skin permeability of drugs. I. Comparison with artificial
ΑU
     Hatanaka, Tomomi; Inuma, Masami; Sugibayashi, Kenji; Morimoto, Yasunori
     Fac. Pharm. Sci., Josai Univ., Sakado, 350-02, Japan
CS
SO
     Chem. Pharm. Bull. (1990), 38(12), 3452-9
     CODEN: CPBTAL; ISSN: 0009-2363
DT
     Journal
LA
     English
SO
     Chem. Pharm. Bull. (1990), 38(12), 3452-9
     CODEN: CPBTAL; ISSN: 0009-2363
ΙT
     Pharmaceutical dosage forms
        (topical, skin permeability of drugs prediction by membrane
        models in relation to)
IT
     50-28-2, .beta.-Estradiol, biological studies
                                                     51-21-8, 5-Fluorouracil
     51-61-6, biological studies
                                 52-31-3, Cyclobarbital
                                                            53-86-1,
     Indomethacin 57-27-2, biological studies
                                                58-15-1, Aminopyrine
     59-92-7, biological studies 60-80-0, Antipyrine 87-33-2, Isosorbide
                                     359-83-1 5104-49-4,
     dinitrate 137-58-6, Lidocaine
     Flurbiprofen
                   7683-59-2
                                15307-86-5
                                           15687-27-1, Ibuprofen
     22071-15-4, Ketoprofen
                              65141-46-0, Nicorandil
                                                       72522-13-5
     RL: PRP (Properties)
        (skin permeability of, membrane models for prediction of)
    ANSWER 11 OF 12 CAPLUS COPYRIGHT 2002 ACS
L10
     Following the topical application of morphine (I)
AΒ
     57-27-2] (0.086-8.6 mM), naloxone (II) [465-65-6] (0.1-1.0 mM),
     or their combination to desheathed saphenous nerves in rats, both drugs
     impaired nerve conduction of single C-fibers in a concn.-dependent but
     reversible fashion. I at 0.86 mM increased the latencies of action
     potential by 10%; when II (0.33 mM) was added to I-treated prepns. nerve
     conduction was decreased by an addnl. 8%, when compared to that obsd. in
     untreated prepns. I and II alter the conduction of peripheral nerves by
     mechanism similar to that obsd. with local anesthetics such as
     lidocaine. A nonopiate action mechanism is postulated. The clin.
     application of I or II for reversible peripheral nerve blockade is not
     recommended since the high doses required for this effect induce the
     well-known opiate side effects.
ΑN
     1986:102377
                 CAPLUS
DN
     104:102377
TI
     Local anesthetic effects of morphine and naloxone
ΑU
     Gilly, H.; Kramer, R.; Zahorovsky, Ingrid
CS
     Exp. Abt. Klin. Anaesth. Allg. Invensivmed., Univ. Wien, Vienna, A-1090,
     Austria
SO
     Anaesthesist (1985), 34(11), 619-26
     CODEN: ANATAE; ISSN: 0003-2417
DT
     Journal
LA
    German
    Local anesthetic effects of morphine and naloxone
TΤ
     Anaesthesist (1985), 34(11), 619-26
SO
    CODEN: ANATAE; ISSN: 0003-2417
    Following the topical application of morphine (I)
AΒ
    57-27-2] (0.086-8.6 mM), naloxone (II) [465-65-6] (0.1-1.0 mM),
    or their combination to desheathed saphenous nerves in rats, both drugs
    impaired nerve conduction of single C-fibers in a concn.-dependent but
    reversible fashion. I at 0.86 mM increased the latencies of action
    potential by 10%; when II (0.33 mM) was added to I-treated prepns. nerve
```

rate of drugs was derived based on this skin permeation model.

```
conduction was decreased by an addnl. 8%, when compared to that obsd. in
     untreated prepns. I and II alter the conduction of peripheral nerves by
а
     mechanism similar to that obsd. with local anesthetics such as
     lidocaine. A nonopiate action mechanism is postulated. The clin.
     application of I or II for reversible peripheral nerve blockade is not
     recommended since the high doses required for this effect induce the
     well-known opiate side effects.
ST
     morphine naloxone local anesthesia; nerve blockade
     morphine naloxone
TΤ
     Neurotransmission
        (blockade of, by morphine and naloxone, local anesthetic
        activity in relation to)
ТТ
     Anesthesia
        (local, from morphine and naloxone)
IT
     57-27-2, biological studies
                                   465-65-6
     RL: BIOL (Biological study)
        (nerve transmission blockade by, local anesthetic activity in relation
        to)
L10
     ANSWER 12 OF 12 CAPLUS COPYRIGHT 2002 ACS
     Improved sepn. of methaqualone (I) [72-44-6], lidocaine [
     137-58-6], cocaine [50-36-2], phencyclidine [77-10-1],
     phenacetin [62-44-2] and methadone [76-99-3] was achieved by a double
     development technique using CH2Cl2:Me Et ketone:NH3 (90:10:1.5 and
     30:70:2). Improved identification of spots for some basic drugs was
     achieved after the traditional detection spray sequence with successive
     topical applications of: Marquis reagent, Mandelin reagent and a
     special NH4 molybdate fuming H2SO4 reagent.
ΑN
     1982:609762 CAPLUS
DN
     97:209762
ΤI
     Improved detection and identification of basic drugs extracted from
tissue
     using TLC
ΑU
     Galante, Lorenzo; Bonventre, Joseph; Salvione, Henry; Bastos, Milton L.
CS
     Inst. Forensic Med., New York, NY, USA
SO
     J. Anal. Toxicol. (1982), 6(5), 262-3
     CODEN: JATOD3; ISSN: 0146-4760
DΤ
     Journal
LA
     English
SO
     J. Anal. Toxicol. (1982), 6(5), 262-3
     CODEN: JATOD3; ISSN: 0146-4760
     Improved sepn. of methaqualone (I) [72-44-6], lidocaine
AB
     137-58-6], cocaine [50-36-2], phencyclidine [77-10-1],
     phenacetin [62-44-2] and methadone [76-99-3] was achieved by a double
     development technique using CH2Cl2:Me Et ketone:NH3 (90:10:1.5 and
     30:70:2). Improved identification of spots for some basic drugs was
     achieved after the traditional detection spray sequence with successive
     topical applications of: Marquis reagent, Mandelin reagent and a
     special NH4 molybdate fuming H2SO4 reagent.
ΙT
               50-47-5
     50-36-2
                         50-48-6
                                   50-53-3, analysis
                                                       57-24-9 57-27-2
     , analysis
                  58-25-3
                            62-44-2
                                      72-44-6
                                                76-57-3
                                                         76-99-3
     91-80-5
             130-95-0 137-58-6
                                   359-83-1
                                            439-14-5
                                                         469-62-5
     1668-19-5
                17617-23-1
     RL: ANT (Analyte); ANST (Analytical study)
        (detn. of, in liver by TLC, forensic)
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